

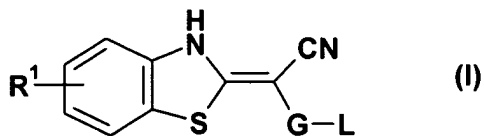
Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

1(Currently amended). A method for treating diabetes type II ~~of decreasing the insulin level in the treatment of a metabolic disorder mediated by insulin resistance or hyperglycemia,~~ comprising administering to a human or other mammal in need thereof:

an effective amount of a compound according to formula I



as well as a tautomer, geometrical isomer, optically active form as enantiomer, diastereomer, racemate, or a pharmaceutically acceptable salt thereof, ~~to decrease the insulin level in the human or other mammal,~~ wherein

G is a pyrimidinyl group;

L is an C₁-C₆-alkoxy, an amino group, or a 3-8 membered heterocycloalkyl, containing at least one

heteroatom selected from the group consisting of N, O, and S; and

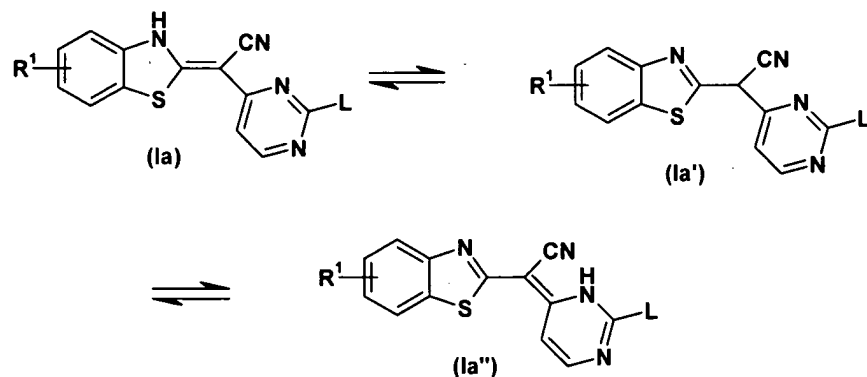
R^1 is selected from the group consisting of hydrogen, sulfonyl, amino, C_1 - C_6 -alkyl, C_2 - C_6 -alkenyl, C_2 - C_6 -alkynyl or C_1 - C_6 -alkoxy, aryl, halogen, cyano and hydroxy; and

at least one supplementary drug selected from the group consisting of insulin, aldose reductase inhibitors, alpha-glucosidase inhibitors, sulfonyl urea agents, biguanides, thiazolidines, PPARs agonists, and GSK-3 inhibitors.

2(Previously presented). The method according to claim 1, wherein the metabolic disorder is diabetes type II.

3(Previously presented). The method according to claim 1, wherein, in the compound, R^1 is H or C_1 - C_3 alkyl.

4(Previously presented). The method according to claim 1, wherein the compound has any of formulae (Ia), (Ia') or (Ia''):



wherein R^1 is selected from the group consisting of hydrogen, sulfonyl, amino, C_1 - C_6 -alkyl, C_2 - C_6 -alkenyl, C_2 - C_6 -alkynyl, C_1 - C_6 -alkoxy, aryl, halogen, cyano, and hydroxy; and

L is an amino group of the formula $-NR^3R^4$, wherein R^3 and R^4 are each independently from each other H, C_1 - C_6 -alkyl, C_2 - C_6 -alkenyl, C_2 - C_6 -alkynyl, C_1 - C_6 -alkoxy, aryl, heteroaryl, saturated or unsaturated 3-8-membered cycloalkyl, 3-8-membered heterocycloalkyl, (wherein said cycloalkyl, heterocycloalkyl, aryl or heteroaryl groups may be fused with 1-2 further cycloalkyl, heterocycloalkyl, aryl or heteroaryl group), C_1 - C_6 -alkyl aryl, C_1 - C_6 -alkyl heteroaryl, C_1 - C_6 -alkenyl aryl, C_1 - C_6 -alkenyl heteroaryl, C_1 - C_6 -alkynyl aryl, C_1 - C_6 -alkynyl heteroaryl, C_1 - C_6 -alkyl cycloalkyl, C_1 - C_6 -alkyl heterocycloalkyl, C_1 - C_6 -alkenyl cycloalkyl, C_1 - C_6 -alkenyl

heterocycloalkyl, C₁-C₆-alkynyl cycloalkyl, C₁-C₆-alkynyl heterocycloalkyl; or

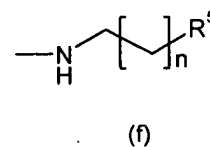
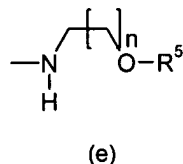
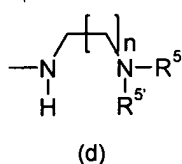
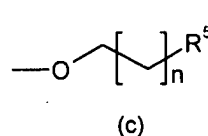
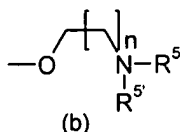
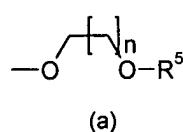
R³ and R⁴ may form a ring together with the nitrogen to which they are bound.

5(Previously presented). The method according to claim 4, wherein, in the compound, R³ is hydrogen or a methyl or ethyl or propyl group and R⁴ is selected from the group consisting of a (C₁-C₆)-alkyl, C₁-C₆-alkyl-aryl, C₁-C₆-alkyl-heteroaryl, cycloalkyl, heterocycloalkyl, aryl or heteroaryl, and 4-8 membered saturated or unsaturated cycloalkyl.

6(Previously presented). The method according to claim 4, wherein, in the compound, R³ and R⁴ form an optionally substituted piperazine or a piperidine or a morpholine or a pyrrolidine ring together with the nitrogen to which they are bound, whereby said optional substituent is selected from the group consisting of a C₁-C₆-alkyl, C₂-C₆-alkenyl, C₂-C₆-alkynyl, C₁-C₆-alkoxy, aryl, heteroaryl, saturated or unsaturated 3-8-membered cycloalkyl, 3-8-membered heterocycloalkyl, (wherein said cycloalkyl, heterocycloalkyl, aryl or heteroaryl groups may be fused with 1-2 further cycloalkyl, heterocycloalkyl, aryl or heteroaryl group), C₁-C₆-alkyl

aryl, C₁-C₆-alkyl heteroaryl, C₁-C₆-alkenyl aryl, C₁-C₆-alkenyl heteroaryl, C₁-C₆-alkynyl aryl, C₁-C₆-alkynyl heteroaryl, C₁-C₆-alkyl cycloalkyl, C₁-C₆-alkyl heterocycloalkyl, C₁-C₆-alkenyl cycloalkyl, C₁-C₆-alkenyl heterocycloalkyl, C₁-C₆-alkynyl cycloalkyl, and C₁-C₆-alkynyl heterocycloalkyl.

7(Previously presented). The method according to claim 5, wherein, in the compound, L is selected from the group consisting of:



wherein n is 1 to 10, and

R⁵ and R^{5'} are independently selected from each other from the group consisting of H, C₁-C₁₀ alkyl, aryl or hetero-aryl, C₁-C₆ alkyl-aryl, and C₁-C₆-alkyl-heteroaryl.

8(Previously presented). The method according to claim 1, wherein the compound is selected from the group consisting of:

Appln. No. 10/571,291
Amd. dated June 16, 2010
Reply to Office Action of March 17, 2010

1,3-benzothiazol-2-yl(2,6-dimethoxy-4-pyrimidinyl)acetonitrile;

1,3-benzothiazol-2-yl(2-{[2-(1H-imidazol-5-yl)ethyl]amino}-4-pyrimidinyl)acetonitrile;

1,3-benzothiazol-2-yl[2-(1-piperazinyl)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl[2-(4-benzyl-1-piperidinyl)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl[2-(4-methyl-1-piperazinyl)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl[2-(4-morpholinyl)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl[2-(methylamino)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl(2-{4-[2-(4-morpholinyl)ethyl]-1-piperazinyl}-4-pyrimidinyl)-acetonitrile;

1,3-benzothiazol-2-yl{2-[4-(benzyloxy)-1-piperidinyl]-4-pyrimidinyl}acetonitrile;

1,3-benzothiazol-2-yl[2-(4-hydroxy-1-piperidinyl)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl(2-{[2-(dimethylamino)ethyl]amino}-4-pyrimidinyl)acetonitrile;

1,3-benzothiazol-2-yl[2-(dimethylamino)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-methoxyethyl)amino]-4-pyrimidinyl}acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-hydroxyethyl)amino]-4-pyrimidinyl}acetonitrile;

1,3-benzothiazol-2-yl[2-(propylamino)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl(2-([3-(1H-imidazol-1-yl)propyl]amino)-4-pyrimidinyl)acetonitrile;

1,3-benzothiazol-2-yl[2-(1-pyrrolidinyl)-4-pyrimidinyl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-phenylethyl)amino]-4-pyrimidinyl}acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(2-pyridinyl)ethyl]amino)-4-pyrimidinyl)acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-pyridinylmethyl)amino]-4-pyrimidinyl}acetonitrile;

1,3-benzothiazol-2-yl{2-[4-(1H-1,2,3-benzotriazol-1-yl)-1-piperidinyl]-4-pyrimidinyl}acetonitrile;

1,3-benzothiazol-2-yl{2-[4-(2-pyrazinyl)-1-piperazinyl]-4-pyrimidinyl}acetonitrile;

Appln. No. 10/571,291
Amd. dated June 16, 2010
Reply to Office Action of March 17, 2010

1,3-benzothiazol-2-yl{2-[4-(2-pyrimidinyl)-1-piperazinyl]-4-pyrimidinyl}acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(3-pyridinyl)ethyl]amino)-4-pyrimidinyl)acetonitrile;

1,3-benzothiazol-2-yl(5-bromo-2-([2-(dimethylamino)ethyl]amino)-4-pyrimidinyl)-acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-morpholin-4-ylethyl)amino]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl[2-(4-{3-[(trifluoromethyl)sulfonyl]anilino}piperidin-1-yl)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl(2-([3-(2-oxopyrrolidin-1-yl)propyl]amino)pyrimidin-4-yl)-acetonitrile;

1,3-benzothiazol-2-yl(2-{methyl[3-(methylamino)propyl]amino}pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([3-(4-methylpiperazin-1-yl)propyl]amino)pyrimidin-4-yl)-acetonitrile;

1,3-benzothiazol-2-yl{2-[(3-morpholin-4-ylpropyl)amino]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(1-methyl-1H-imidazol-4-yl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(1H-indol-3-yl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(4-hydroxyphenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

tert-butyl ({4-[1,3-benzothiazol-2-yl(cyano)methyl]pyrimidin-2-yl}amino)acetate

(2-[(3-aminopropyl)amino]pyrimidin-4-yl)(1,3-benzothiazol-2-yl)acetonitrile;

(2-[(2-aminoethyl)amino]pyrimidin-4-yl)(1,3-benzothiazol-2-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([3-(dimethylamino)propyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-piperidin-1-ylethyl)amino]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(1-methyl-1H-imidazol-5-yl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl[2-(benzylamino)pyrimidin-4-yl]acetonitrile;

isopropyl 3-([4-[1,3-benzothiazol-2-yl(cyano)methyl]pyrimidin-2-yl]amino)propanoate;

1,3-benzothiazol-2-yl{2-[(3-hydroxypropyl)amino]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl{2-[(pyridin-3-ylmethyl)amino]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl{2-[(pyridin-4-ylmethyl)amino]pyrimidin-4-yl}acetonitrile;

tert-butyl 4-[2-({4-[1,3-benzothiazol-2-yl(cyano)methyl]pyrimidin-2-yl}amino)-ethyl]phenylcarbamate;

(2-{{2-(4-aminophenyl)ethyl}amino}pyrimidin-4-yl)(1,3-benzothiazol-2-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-{{2-(3,4-dimethoxyphenyl)ethyl}amino}pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-{{2-(3-methoxyphenyl)ethyl}amino}pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-{{2-(2-fluorophenyl)ethyl}amino}pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl[2-({2-[3-(trifluoromethyl)phenyl]ethyl}amino)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-hydroxy-2-phenylethyl)amino]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl{2-[(2-{{3-(trifluoromethyl)pyridin-2-yl}amino)ethyl}amino]-pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl(2-{{2-(3-chlorophenyl)ethyl}amino}pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(3,4-dichlorophenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(4-methoxyphenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(4-methylphenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(3-fluorophenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(4-phenoxyphenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(2-phenoxyphenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(4-bromophenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(4-fluorophenyl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-([1,1'-biphenyl]-4-ylethyl)amino]pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(4-[hydroxy(oxido)amino]phenyl)ethyl]amino]pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([2-(1H-1,2,4-triazol-1-yl)ethyl]amino)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl(2-([3-(1H-pyrazol-1-yl)propyl]amino)pyrimidin-4-yl)acetonitrile;

4-[2-([4-[1,3-benzothiazol-2-yl(cyano)methyl]pyrimidin-2-yl]amino)ethyl]benzenesulfonamide;

{2-[(2-pyridin-3-ylethyl)amino]pyrimidin-4-yl}[5-(trifluoromethyl)-1,3-benzothiazol-2-yl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(1H-tetraazol-5-ylmethyl)amino]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl[2-(benzyloxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(4-pyridin-3-ylbenzyl)oxy]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl[2-(pyridin-4-ylmethoxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl[2-(pyridin-2-ylmethoxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl[2-(3-pyridin-2-ylpropoxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(4-methoxybenzyl)oxy]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl[2-(pyridin-3-ylmethoxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl{2-[2-(4-methoxyphenyl)ethoxy]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl[2-([1,1'-biphenyl]-3-ylmethoxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(3,4,5-trimethoxybenzyl)oxy]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl{2-[(3,4-dichlorobenzyl)oxy]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl[2-({3-[(dimethylamino)methyl]benzyl}oxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl{2-[(1-oxidopyridin-3-yl)methoxy]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl(2-{[4-(morpholin-4-ylmethyl)benzyl]oxy}pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl{2-[(4-pyridin-2-ylbenzyl)oxy]pyrimidin-4-yl}acetonitrile;

1,3-benzothiazol-2-yl(2-{[4-(piperidin-1-ylmethyl)benzyl]oxy}pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl[2-(4-methoxyphenoxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl[2-(4-butoxyphenoxy)pyrimidin-4-yl]acetonitrile;

{2-[4-(4-acetylpiperazin-1-yl)phenoxy]pyrimidin-4-yl}(1,3-benzothiazol-2-yl)acetonitrile;

[2-(4-methoxyphenoxy)pyrimidin-4-yl][5-(trifluoromethyl)-1,3-benzothiazol-2-yl]acetonitrile;

N-[2-({4-[1,3-benzothiazol-2-yl(cyano)methyl]pyrimidin-2-yl}amino)ethyl)-4-chlorobenzamide;

1,3-benzothiazol-2-yl(2-methoxy-4-pyrimidinyl)acetonitrile;

1,3-benzothiazol-2-yl[2-({4-[(4-methylpiperazin-1-yl)methyl]benzyl}oxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl[2-({4-[(4-benzyl-piperazin-1-yl)methyl]-benzyl}oxy)pyrimidin-4-yl]acetonitrile;

1,3-benzothiazol-2-yl(2-({4-(piperazin-1-ylmethyl)benzyl}oxy)pyrimidin-4-yl)acetonitrile;

1,3-benzothiazol-2-yl[2-({4-[(4-formylpiperazin-1-yl)methyl]benzyl}oxy)pyrimidin-4-yl]acetonitrile;

[2-({4-[(4-acetylpiperazin-1-yl)methyl]benzyl}oxy)pyrimidin-4-yl](1,3-benzothiazol-2-yl)acetonitrile;

(3H-Benzothiazol-2-ylidene)-(2-[4-(4-[1,2,4]oxadiazol-3-ylmethyl-piperazin-1-ylmethyl)-benzyloxy]-pyrimidin-4-yl)-acetonitrile;

4-(4-(4-[(3H-Benzothiazol-2-ylidene)-cyano-methyl]-pyrimidin-2-yloxymethyl)-benzyl)-piperazine-1-carboxylic acid methyl ester;

2-[4-(4-(4-[(3H-Benzothiazol-2-ylidene)-cyano-methyl]-pyrimidin-2-yloxymethyl)-benzyl)-piperazin-1-yl]-acetamide;

(2-{4-[4-(2-Amino-acetyl)-piperazin-1-ylmethyl]-benzyloxy}-pyrimidin-4-yl)-(3H-benzothiazol-2-ylidene)-acetonitrile;

[4-(4-(4-[(3H-Benzothiazol-2-ylidene)-cyano-methyl]-pyrimidin-2-yloxymethyl)-benzyl)-piperazin-1-yl]-acetic acid methyl ester;

(3H-Benzothiazol-2-ylidene)-(2-{4-[4-(2-methoxy-ethyl)-piperazin-1-ylmethyl]-benzyloxy}-pyrimidin-4-yl)-acetonitrile;

4-(4-(4-[(3H-Benzothiazol-2-ylidene)-cyano-methyl]-pyrimidin-2-yloxymethyl)-benzyl)-piperazine-1-carboxylic acid dimethylamide;

(3H-Benzothiazol-2-ylidene)-{2-[4-(4-ethyl-piperazin-1-ylmethyl)-benzyloxy]-pyrimidin-4-yl}-acetonitrile; and

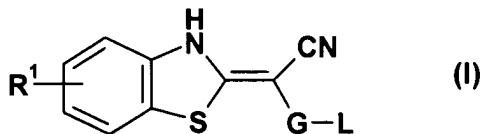
(3H-Benzothiazol-2-ylidene)-(2-{4-[4-(2-hydroxy-ethyl)-piperazin-1-ylmethyl]-benzyloxy}-pyrimidin-4-yl)-acetonitrile.

Claim 9 (Cancelled).

10(Currently amended). The method according to claim 91, wherein said supplementary drug is selected from the group consisting of a rapid acting insulin, an intermediate acting insulin, a long acting insulin, a combination of intermediate and rapid acting insulins, Minalrestat, Tolrestat, Sorbinil, Methosorbinil, Zopolrestat, Epalrestat, Zenarestat, Imirestat, Ponalrestat, ONO-2235, GP-1447, CT-112, BAL-ARI 8, AD-5467, ZD5522, M-16209, NZ-314, M-79175, SPR-210, ADN 138, or SNK-860, Miglitol, Acarbose, Glipizide, Glyburide, Chlorpropamide, Tolbutamide, Tolazamide, and Glimepiride.

11(Previously presented). The method according to claim 1, wherein n is 1 to 6.

12(Previously presented). A pharmaceutical composition comprising an anti-diabetes agent and a compound according to formula I:



as well as a tautomer, geometrical isomer, optically active form as enantiomer, diastereomer, racemate, or a pharmaceutically acceptable salt thereof, wherein

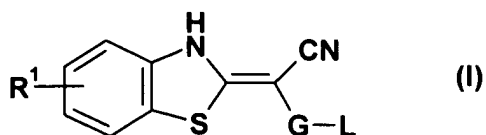
G is a pyrimidinyl group;

L is an C₁-C₆-alkoxy, an amino group, or a 3-8 membered heterocycloalkyl, containing at least one heteroatom selected from the group consisting of N, O, and S; and

R¹ is selected from the group consisting of hydrogen, sulfonyl, amino, C₁-C₆-alkyl, C₂-C₆-alkenyl, C₂-C₆-alkynyl or C₁-C₆-alkoxy, aryl, halogen, cyano and hydroxy.

13(Currently amended). A method for treating diabetes II ~~of decreasing the insulin level in the treatment of a metabolic disorder mediated by insulin resistance or hyperglycemia,~~ comprising administering an effective amount of the pharmaceutical composition according to claim 12 to a human or other mammal in need thereof ~~to decrease the insulin level in the human or other mammal.~~

14(Currently amended). A method for the preparation of a pharmaceutical composition for treating diabetes II ~~decreasing the insulin level in the treatment of metabolic disorders mediated by insulin resistance or hyperglycemia,~~ comprising combining a compound with an anti-diabetes agent, wherein the compound is one according to formula I:



Appln. No. 10/571,291
Amd. dated June 16, 2010
Reply to Office Action of March 17, 2010

as well as a tautomer, geometrical isomer,
optically active form as enantiomer, diastereomer,
racemate, or a pharmaceutically acceptable salt thereof,
wherein

G is a pyrimidinyl group;

L is an C₁-C₆-alkoxy, an amino group, or a 3-8
membered heterocycloalkyl, containing at least one heteroatom
selected from the group consisting of N, O, and S; and

R¹ is selected from the group consisting of hydrogen,
sulfonyl, amino, C₁-C₆-alkyl, C₂-C₆-alkenyl, C₂-C₆-alkynyl or C₁-
C₆-alkoxy, aryl, halogen, cyano and hydroxy.

Claims 15-20 (Cancelled).